

GenCore version 4.5  
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OM protein - protein search, using sw model

Run on: March 15, 2001, 10:36:10 ; Search time 35.6 Seconds  
(without alignments)  
4.802 Million cell updates/sec

Title: US-09-288-719-1  
Perfect score: 28  
Sequence: 1 GCGGS 5

Scoring table: BLOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 268485 seqs, 34193795 residues  
Total number of hits satisfying chosen parameters: 268485

Minimum DB seq length: 0  
Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%  
Maximum Match 100%  
Listing first 45 summaries

Database :

A\_Geneseq\_36:\*

- 1: /SIDSL/gcgdata/geneseq/geneseqp/AA1980.DAT:\*
- 2: /SIDSL/gcgdata/geneseq/geneseqp/AA1981.DAT:\*
- 3: /SIDSL/gcgdata/geneseq/geneseqp/AA1982.DAT:\*
- 4: /SIDSL/gcgdata/geneseq/geneseqp/AA1983.DAT:\*
- 5: /SIDSL/gcgdata/geneseq/geneseqp/AA1984.DAT:\*
- 6: /SIDSL/gcgdata/geneseq/geneseqp/AA1985.DAT:\*
- 7: /SIDSL/gcgdata/geneseq/geneseqp/AA1986.DAT:\*
- 8: /SIDSL/gcgdata/geneseq/geneseqp/AA1987.DAT:\*
- 9: /SIDSL/gcgdata/geneseq/geneseqp/AA1988.DAT:\*
- 10: /SIDSL/gcgdata/geneseq/geneseqp/AA1989.DAT:\*
- 11: /SIDSL/gcgdata/geneseq/geneseqp/AA1990.DAT:\*
- 12: /SIDSL/gcgdata/geneseq/geneseqp/AA1991.DAT:\*
- 13: /SIDSL/gcgdata/geneseq/geneseqp/AA1992.DAT:\*
- 14: /SIDSL/gcgdata/geneseq/geneseqp/AA1993.DAT:\*
- 15: /SIDSL/gcgdata/geneseq/geneseqp/AA1994.DAT:\*
- 16: /SIDSL/gcgdata/geneseq/geneseqp/AA1995.DAT:\*
- 17: /SIDSL/gcgdata/geneseq/geneseqp/AA1996.DAT:\*
- 18: /SIDSL/gcgdata/geneseq/geneseqp/AA1997.DAT:\*
- 19: /SIDSL/gcgdata/geneseq/geneseqp/AA1998.DAT:\*
- 20: /SIDSL/gcgdata/geneseq/geneseqp/AA1999.DAT:\*
- 21: /SIDSL/gcgdata/geneseq/geneseqp/AA2000.DAT:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

#### SUMMARIES

Result No.	Score	Query Match	Length	ID	Description
1	28	100.0	5	14 R34034	Linking sequence w
2	28	100.0	5	16 R72707	Linker for apo A-I
3	28	100.0	5	17 R95062	scfv spacer peptid
4	28	100.0	5	18 W17094	GLY(4)-Ser linker
5	28	100.0	5	18 W19543	Chimeric protein p
6	28	100.0	5	20 Y43496	Linker for dual av
7	28	100.0	5	20 Y33597	VH-VL domain linke
8	28	100.0	5	20 Y25357	IFNAR2/IFN-beta co
9	28	100.0	5	20 Y02127	Peptide linker use
10	28	100.0	5	21 Y83210	Peptide linker use
11	28	100.0	5	21 Y43750	Linker used to con
12	28	100.0	5	21 Y54917	Linker from IL-12

13	28	100.0	6	15 R62168	U1 snRNP 70K prote
14	28	100.0	6	18 W17095	GLY(5)-Ser linker
15	28	100.0	6	18 W21967	Linker #1 for immu
16	28	100.0	7	17 R99245	(GLY6)Ser linker..
17	28	100.0	7	20 Y23703	Peptide identified
18	28	100.0	7	20 Y02129	Peptide linker use
19	28	100.0	8	17 R86795	GM-CSF/EPD linker
20	28	100.0	8	20 Y43498	Linker for dual av
21	28	100.0	8	21 Y83212	Peptide linker use
22	28	100.0	9	14 R31941	In vivo tumour bin
23	28	100.0	9	16 R77978	Conserved TDP2 pep
24	28	100.0	9	18 W08976	Conserved epitope
25	28	100.0	9	18 W43015	Conserved epitope
26	28	100.0	9	19 W54140	H. influenzae TBP2
27	28	100.0	9	19 W54141	H. influenzae TBP2
28	28	100.0	9	21 Y51796	H. influenzae tran
29	28	100.0	9	21 Y51797	H. influenzae tran
30	28	100.0	9	21 Y80384	H. influenzae tran
31	28	100.0	9	21 Y80385	H. influenzae tran
32	28	100.0	11	17 R99242	(GLY4Ser)2Ser link
33	28	100.0	11	17 R91061	Linker peptide use
34	28	100.0	11	19 W59848	Amino acid sequenc
35	28	100.0	12	21 Y79553	Linker peptide use
36	28	100.0	13	20 Y43499	Linker for dual av
37	28	100.0	13	20 Y25363	IFNAR2/IFN-beta co
38	28	100.0	13	20 Y06843	Peptide sequence f
39	28	100.0	13	21 Y83213	Peptide linker use
40	28	100.0	13	21 Y83220	Peptide linker use
41	28	100.0	13	21 Y80115	IL-6R and IL-6 fus
42	28	100.0	13	21 Y44696	Peptide linker to
43	28	100.0	14	16 R87024	Flexible linker se
44	28	100.0	14	18 W23417	Linker peptide for
45	28	100.0	14	19 W64498	Neurotoxic Beta-am

#### ALIGNMENTS

RESULT 1

R34034 R34034 standard; Protein: 5 AA.

AC R34034;

DT 13-AUG-1993 (first entry)

DE Linking sequence whose encoding DNA can be ligated between an

apo A-I - and a B-100-encoding DNA sequence.

KM Lipoprotein; apoprotein; B-100; A-I; LDL; HDL; assay.

OS Synthetic.

PN W09307165-A.

PD 15-APR-1993.

PF 09-OCT-1992; 92WO-US08634.

PR 09-OCT-1991; 91US-0774633.

PR 08-OCT-1992; 92US-0555555.

PR 28-JUN-1992; 92US-0901706.

PA (SCRI ) SCRIPPS RES INSTR.

PI Curtiss LK, Koduri KR, Smith RS, Wiltzum JL, Young SG;

DR WPI; 1993-134378/16.

PT Polypeptide mimic of native apo B-100 and native apo A-I - useful

PS in assays for LDL and HDL in plasma samples  
Disclosure: Page 14 and page 35; 137pp; English.

XX The inventors claim a portion of the polypeptide contg. apo B-100  
CC that immunoreacts with antibodies secreted by the hybridoma MB47  
CC having ATCC Accession No. 8746. Polypeptides specifically claimed  
CC include residues 217-287, 216-310, 216-331, 216-352, 216-377, 1-377,  
CC 205-297, 173-297, 140-297. DNA sequences encoding the polypeptides  
CC are also claimed. Also claimed are a fusion polypeptide that  
CC contains: (a) a first amino acid residue sequence up to 250 residues  
CC in length that includes residues 120-135 of apo A-I, (b) a second  
CC amino acid residue sequence up to 375 residues in length that  
CC includes residues 217-297 of apo B-100 and DNA encoding it.  
XX  
SQ Sequence 5 AA:

Query Match 100.0%; Score 28; DB 14; Length 5;  
Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

OY 1 GGGGS 5  
| | | | |  
Db 1 ggggs 5

RESULT 2  
R72707  
ID R72707 standard; Peptide; 5 AA.  
XX  
AC R72707;  
XX  
DT 31-OCT-1995 (first entry)  
XX

DE Linker for apo A-I and apo B-100 fusion polypeptide.

XX Apo A-I; LDL cholesterol; low density lipoprotein;  
KM fusion polypeptide; linker.  
XX

OS Synthetic.

XX  
PN US5408038-A.  
XX

PD 18-APR-1995.  
XX

PF 09-OCT-1991; 91US-0774633.  
XX

PR 09-OCT-1991; 91US-0774633.  
XX

PR 18-JUN-1992; 92US-0901706.  
XX

PR 08-OCT-1992; 92US-0959946.  
XX

PA (SCRI ) SCRIPPS RES INST.  
XX

PI Curtiss IK, Koduri KR, Smith RS, Witzum JL, Young SG;  
XX

DR WPI: 1995-161146/21.  
XX

PT New apo:lipoprotein B-100 peptide(s) and fusion peptide(s) - used  
XX in assay systems for detecting LDL and HDL cholesterol levels in  
XX body fluids.  
XX

PS Disclosure; Column 18; 41pp; English.

XX A dispersible apo A-I/B-100 fusion polypeptide is claimed which  
CC contains a first AA sequence of apo A-I (see R72605) and that includes  
CC at least AA sequence positions 120-135 (see R72606). The two  
CC sequences are operatively linked. An exemplary linking sequence is  
CC R72707 whose encoding DNA can be ligated between an apo A-I and a  
CC B-100 encoding DNA sequence.  
XX

SQ Sequence 5 AA:

Query Match 100.0%; Score 28; DB 16; Length 5;  
Best Local Similarity 100.0%; Pred. No. 2.1e+05;

Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
OY 1 GGGGS 5  
| | | | |  
Db 1 ggggs 5

RESULT 3  
R95062  
ID R95062 standard; Peptide; 5 AA.  
XX  
AC R95062;  
XX

DT 18-AUG-1996 (first entry)  
XX

DE scfv spacer peptide.

XX Nucleic acid transfer system; gene transfer; gene therapy;  
KM cell targeting; multidomain protein; vector; cancer; scfv;  
KW single chain antibody.  
XX

OS Synthetic.

XX WO9613599-A1.  
XX

PD 09-MAY-1996.  
XX

PF 31-OCT-1995; 95WO-EP04270.  
XX

PR 01-NOV-1994; 94EP-0810627.  
XX

PA (WELS/) WELS W.  
XX

PI Fominaya J, Wels W;  
XX

DR WPI: 1996-239505/24.  
XX

PT Nucleic acid transfer system for gene therapy, e.g. against cancer  
XX - includes toxin translocation domain to target nucleic acid to  
XX specific cell  
XX

PS Disclosure; Page 8; 106pp; English.

CC A flexible spacer peptide (R95062) is used to link the light chain  
CC variable domain to the heavy chain variable domain of a single  
CC chain recombinant antibody (scfv). The scfv may be derived from  
CC a monoclonal antibody, e.g. Mab FRP5, and forms the ligand domain  
CC of a multidomain protein (see also R95053 and R95056-58) that is used  
CC with an effector nucleic acid in a novel nucleic acid transfer system  
CC suitable for gene therapy. The ligand domain has a target cell  
CC recognition function and allows cellular internalization of the  
CC multidomain protein/nucleic acid complex.  
XX

SQ Sequence 5 AA:

Query Match 100.0%; Score 28; DB 17; Length 5;  
Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

OY 1 GGGGS 5  
| | | | |  
Db 1 ggggs 5

RESULT 4  
W17094  
ID W17094 standard; peptide; 5 AA.  
XX  
AC W17094;  
XX

DT 14-SEP-1999 (first entry)  
XX

DE Gly(4)-Ser linker peptide for chimeric protein construct.  
 XX  
 KW Haematopoietic protein; human; granulocyte-colony stimulating factor;  
 KW G-CSF; interleukin; c-mpl ligand; linker; gene therapy; aplastic anaemia;  
 KW stem cell expansion; leukaemia; neutropenia; vector; bone marrow;  
 KW thrombocytopaenia; blood cell activation; growth.  
 XX  
 OS Synthetic.  
 XX  
 PN WO9712985-A2.  
 XX  
 PD 10-APR-1997.  
 XX  
 PF 04-OCT-1996; 96WO-US15774.  
 XX  
 PR 05-OCT-1995; 95US-0004834.  
 XX  
 PA (SEAR ) SEARLE & CO G. D.  
 XX  
 PI Bauer SC, Baum CM, Caparon MH, Feng Y, Gird JG;  
 PI Klein BK, Lee SC, McKearn JP, McWhorter CA, Staten NR;  
 PI Summers NL, Zurfluh L;  
 XX  
 DR WPI; 1997-226228/20.  
 XX  
 PT Multi-functional haematopoietic receptor agonists - used to  
 PT stimulate the production of haematopoietic cells in patients  
 XX  
 PS Disclosure; Page 33; 616pp; English.  
 XX  
 CC The invention relates to a novel haematopoietic protein (HP) comprising  
 CC an amino acid (AA) sequence of formula: R1-L1-R2; R2-L1-R1; R1-R2; or  
 CC R2-R1; where R1 and R2 are independently selected from: (i) a modified  
 CC human granulocyte-colony stimulating factor (hG-CSF) AA sequence;  
 CC (ii) a modified human interleukin-3 (hIL-3) AA sequence; (iii) a modified  
 CC human c-mpl ligand; and a colony stimulating factor (CSF); and L1 = a  
 CC linker capable of linking R1 to R2. This sequence represents an example  
 CC of a linker used to construct the proteins of the invention.  
 CC Vectors comprising the nucleic acid molecules are useful for the  
 CC recombinant production of HP. The nucleic acid molecules are useful in  
 CC gene therapy. The HP's are useful for stimulating the production of  
 CC haematopoietic cells in patients, selective ex vivo expansion of stem  
 CC cells and for treatment of haematopoietic disorders. Disorders that  
 CC can be treated include leukaemia, neutropenia, aplastic anaemia and  
 CC thrombocytopaenia. In vitro uses include the ability to stimulate bone  
 CC marrow and blood cell activation and growth before infusion into the  
 CC patients.  
 CC  
 CC Sequence 5 AA;  
 SQ

Query Match 100.0%; Score 28; DB 18; Length 5;  
 Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
 Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

OY 1 GGGGS 5  
 |||||  
 DB 1 9999s 5

RESULT 5  
 W19543  
 ID W19543 standard; peptide; 5 AA.  
 XX  
 AC W19543;  
 XX  
 DT 19-FEB-1998 (first entry)  
 XX  
 DE Chimeric protein pentapeptide linker for the MBP moiety and PE moiety.  
 XX  
 KW Pseudomonas exotoxin; myelin basic protein; chimeric protein;  
 KW autoimmune disease; multiple sclerosis; human.  
 XX  
 OS Synthetic.

XX  
 PN WO9719179-A1.  
 XX  
 PD 29-MAY-1997.  
 XX  
 PF 17-NOV-1996; 96WO-IL00151.  
 XX  
 PR 26-DEC-1995; 95IL-0116559.  
 XX  
 PR 17-NOV-1995; 95IL-0116044.  
 XX  
 PA (YISS ) YISSUM RES & DEV CO.  
 XX  
 PI Beraud E, Lorberboun-Galski H, Marianovsky I, Steinberger I;  
 PI Yarkoni S;  
 XX  
 DR WPI; 1997-298116/27.  
 XX  
 PT New Pseudomonas exotoxin-myelin basic protein chimeric proteins -  
 PT used for the treatment of autoimmune diseases, particularly  
 PT multiple sclerosis  
 XX  
 PS Claim 6; Page 29; 54pp; English.  
 XX  
 CC New chimeric proteins have been developed comprising a Pseudomonas  
 CC aeruginosa exotoxin (PE) moiety linked to a myelin basic protein (MBP)  
 CC moiety selected from: (a) MBP; (b) amino acids 69-88 of guinea-pig MBP  
 CC or an antigenic portion; (c) amino acids 84-102 of human MBP or an  
 CC antigenic portion; (d) amino acids 143-168 of human MBP or an antigenic  
 CC portion; and (e) an amino acid sequence in which one or more amino acids  
 CC have been deleted, added, substituted or mutated in the amino acid  
 CC sequences of (a), (b), (c), or (d), the modified sequences retaining at  
 CC least 75% homology with the amino acid sequences. The present sequence  
 CC represents the preferred pentapeptide linker used to link the MBP moiety  
 CC and PE moiety in a chimeric protein. The chimeric proteins can be used  
 CC for the treatment of autoimmune diseases such as multiple sclerosis. The  
 CC chimeric proteins can specifically target and kill MBP specific T cells  
 CC while having no effect on non-target cells.  
 CC  
 CC Sequence 5 AA;  
 SQ

Query Match 100.0%; Score 28; DB 18; Length 5;  
 Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
 Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

OY 1 GGGGS 5  
 |||||  
 DB 1 9999s 5

RESULT 6  
 Y43496  
 ID Y43496 standard; Peptide; 5 AA.  
 XX  
 AC Y43496;  
 XX  
 DT 26-JAN-2000 (first entry)  
 XX  
 DE Linker for dual avb3 receptor/metastasis-associated receptor ligands.  
 XX  
 KW Interferon-alpha-2b; IFN-alpha; avb3 antagonist; avb3 receptor ligand;  
 KW metastasis-associated receptor ligand; angiogenesis; cell proliferation;  
 KW anti-angiogenic protein; avb3-integrin; cancer; arthritis;  
 KW macular degeneration; diabetic retinopathy; hemangioma; psoriasis;  
 KW osteoporosis; thrombosis; angina; atherosclerosis; antiviral;  
 KW antibacterial; antifungal.  
 XX  
 OS Homo sapiens.  
 XX  
 PN WO9951638-A1.  
 XX  
 PD 14-OCT-1999.

PF 07-APR-1999; 99WO-US04295.  
 XX  
 XX 08-APR-1998; 98US-0081074.  
 XX  
 PA (SEAR ) SEARLE & CO G. D.  
 XX  
 PI Tjoeng FS, Fok KF;  
 XX  
 DR WPI: 1999-620196/53.  
 XX  
 PT New conjugates of integrin antagonist and ligand for  
 PT metastasis-associated receptor, for treating angiogenesis-related  
 PT diseases, e.g. cancer  
 XX  
 PS Claim 18; Page 86; 108pp; English.  
 XX  
 CC The present sequence represents a linker used to join the avb3  
 CC antagonist and the metastasis-associated receptor ligand, in the  
 CC pharmaceutical compounds of the invention. These compounds are dual  
 CC avb3 receptor/metastasis-associated receptor ligands, and inhibit  
 CC angiogenesis and thus proliferation of (cancer) cells. One component  
 CC binds to the avb3 receptor and the other to a metastasis-associated  
 CC receptor. The avb3 antagonists may also be conjugated to anti-angiogenic  
 CC proteins, such as IFN-alpha and its derivatives. The compounds are used  
 CC to treat angiogenesis-related disorders (mediated by the avb3-integrin),  
 CC specifically cancer (of lung, breast, prostate, stomach, colon,  
 CC kidney or bladder, also melanoma, hepatoma, sarcoma and lymphoma),  
 CC arthritis and macular degeneration, and also diabetic retinopathy,  
 CC hemangioma, psoriasis, osteoporosis, thrombosis, angina, atherosclerosis  
 CC etc. The compounds may also be useful as antiviral, antibacterial and  
 CC antifungal agents.  
 CC  
 CC Sequence 5 AA;  
 CC  
 CC SO

Query Match 100.0%; Score 28; DB 20; Length 5;  
 Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
 Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

OY 1 GGGGS 5  
 11111  
 Db 1 9999s 5

RESULT 7  
 Y33597  
 ID Y33597 standard; Protein: 5 AA.  
 XX  
 AC Y33597;  
 XX  
 DT 20-DEC-1999 (first entry)  
 XX  
 DE VH-VL domain linker peptide #9.  
 XX  
 DE heavy chain; variable domain; VH domain; light chain;  
 XX antigen binding; single chain; VL domain; anticancer; antiviral; tumor;  
 XX antibacterial; antimalarial; antiinflammatory; treatment; prevention;  
 XX diagnosis; vaccine; autoimmune disease; inflammation; blood disorder;  
 XX transplant rejection; arthritis; nervous system disorder; infection.  
 XX  
 OS Synthetic.  
 XX  
 PN DE19816141-A1.  
 XX  
 PD 14-OCT-1999.  
 XX  
 PF 09-APR-1998; 98DE-1016141.  
 XX  
 PR 09-APR-1998; 98DE-1016141.  
 XX  
 PA (HMRI ) HOECHST MARION ROUSSEL DEUT GMBH.  
 XX  
 PI Konfermann R, Sedlacek H, Mueller R;  
 XX

XX  
 DR WPI: 1999-581511/50.  
 XX  
 XX New polyspecific binding agents containing variable heavy and light  
 PT constructs connected via peptide linker, used for treatment, prevention  
 PT or diagnosis of e.g. cancer  
 XX  
 PS Claim 7; Page 17; 20pp; German.  
 XX  
 CC This sequence represents a novel single-chain molecule (I) that binds  
 CC multiple antigens and comprises two variable domains of heavy  
 CC immunoglobulin chains (VH), having specificities A and B and two  
 CC variable domains of light chains (VL), also with specificities A and B.  
 CC The domains are provided as two VH-VL constructs which are attached via  
 CC a peptide (P). Any VH and VL may be replaced by their functional  
 CC fragments. The products of the invention have anticancer, antiviral,  
 CC antibacterial, antimalarial and antiinflammatory activity. (I) are used  
 CC to treat, prevent or diagnose tumors (e.g. as tumor vaccines), autoimmune  
 CC diseases and inflammation (e.g. transplant rejection and arthritis),  
 CC blood disorders (e.g. of the coagulation and/or circulatory systems, such  
 CC as anemia, leucopenia, thrombocytopenia and hypertension), nervous system  
 CC disorders and/or infections (by viruses or bacteria, or malaria),  
 CC including, when (I) include a fusogenic peptide, use for gene transfer.  
 CC (I) are produced simply and in predominantly homogeneous form, in a wide  
 CC variety of hosts, either in secreted or membrane-bound forms. This  
 CC sequence represents a VH-VL domain linker peptide which is used to  
 CC illustrate the method of the invention.  
 CC  
 CC Sequence 5 AA;  
 CC  
 CC SO

Query Match 100.0%; Score 28; DB 20; Length 5;  
 Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
 Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

OY 1 GGGGS 5  
 11111  
 Db 1 9999s 5

RESULT 8  
 Y25357  
 ID Y25357 standard; peptide: 5 AA.  
 XX  
 AC Y25357;  
 XX  
 DT 06-SEP-1999 (first entry)  
 XX  
 DE IFNAR2/IFN-beta complex peptide fragment 1.  
 XX  
 DE IFNAR2: IFN-beta; type I interferon; IFNAR/IFN complex; IFN; antiviral;  
 XX human interferon alpha/beta receptor; anticancer; immunomodulatory;  
 XX anti-arthritis; antidiabetic; treatment; hepatitis; viral infection;  
 XX hairy cell leukemia; Kaposi's sarcoma; multiple myeloma; cancer; lupus;  
 XX diabetes; multiple sclerosis; rheumatoid arthritis; myasthenia gravis;  
 XX acquired immune deficiency syndrome.  
 XX  
 OS Synthetic.  
 XX  
 PN WO9932141-A1.  
 XX  
 PD 01-JUL-1999.  
 XX  
 PF 18-DEC-1998; 98WO-US26926.  
 XX  
 PR 19-DEC-1997; 97US-0068295.  
 XX  
 PA (ISTF ) ARS APPLIED RES SYSTEMS HOLDING NV.  
 XX (MCIN/) MCINNIS P G.  
 XX  
 PI Cunningham M, El Tayar N, McKenna S, Sherris D;  
 XX  
 PI Tepper M;  
 XX

DR WPI: 1999-405115/34.  
 XX Prolonging in vivo activity of type I interferon by complexing  
 XX  
 XX  
 PS Example 8; Page 76; 86pp; English.  
 XX  
 CC This invention describes a novel method for prolonging the in vivo effect  
 CC of type I interferon (IFN) by administering IFN as a complex (A) with a  
 CC subunit (I) of the human interferon alpha/beta receptor (IFNAR). The  
 CC product of the invention has antiviral, anticancer, immunomodulatory,  
 CC anti-arthritic and antidiabetic activity. (A) have the antiviral,  
 CC anticancer and immunomodulating activities of IFN, e.g. for treating  
 CC hepatitis and other viral infections, hairy cell leukemia, Kaposi's  
 CC sarcoma, multiple myeloma and other cancers, multiple sclerosis,  
 CC rheumatoid arthritis, myasthenia gravis, diabetes, acquired immune  
 CC deficiency syndrome and lupus. When complexed in (A), the storage life of  
 CC IFN is increased (i.e. it is stabilized against oligomerization, without  
 CC the need for storage at acidic pH) and its biological effect is  
 CC potentiated.  
 CC  
 CC Sequence 5 AA;  
 SO  
 Query Match 100.0%; Score 28; DB 20; Length 5;  
 Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
 Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 QY 1 GGGGS 5  
 Db 1 11111  
 1 999gs 5  
 RESULT 9  
 ID Y02127 standard; Protein; 5 AA.  
 XX Y02127;  
 AC  
 XX 16-JUL-1999 (first entry)  
 DT  
 XX Peptide linker used to make multifunctional proteins.  
 DE  
 XX Angiostatin; endostatin; interferon; thrombospondin;  
 KW interferon-inducible protein; platelet factor 4; anti-angiogenic;  
 KW anti-tumor; multifunctional protein; angiogenic-mediated disease;  
 KW cancer; diabetic retinopathy; macular degeneration; arthritis;  
 KW tumor cell production; peptide linker.  
 KW  
 XX Homo sapiens.  
 OS  
 XX WO9916889-A1.  
 PN  
 XX 08-APR-1999.  
 PD  
 XX 30-SEP-1998; 98WO-US20464.  
 PE  
 XX 01-OCT-1997; 97US-0060609.  
 PR  
 XX (SEAR ) SEARLE & CO G D.  
 PA  
 XX Bolanowski MA, Caparon MH, Casperson GF, Gregory SA;  
 PI Klein BK, McKearn JP;  
 DR WPI: 1999-255098/21.  
 XX  
 XX New multifunctional proteins useful for treating angiogenic-mediated  
 PT diseases  
 PS Disclosure; Page 111; 121pp; English.  
 XX  
 CC The specification describes multifunctional proteins which comprise  
 CC combinations of angiostatin, endostatin, interferon, thrombospondin,  
 CC interferon-inducible protein and platelet factor 4, and have

CC anti-angiogenic and/or anti-tumor activity. The multifunctional protein  
 CC may exhibit useful properties such as having similar or greater  
 CC biological activity when compared to a single factor or by having  
 CC improved half-life or decreased adverse side effects, or a combination  
 CC of these properties. The proteins can be used for treating an  
 CC angiogenic-mediated disease, e.g. cancer, diabetic retinopathy, macular  
 CC degeneration, or arthritis. They can also be used for inhibiting the  
 CC production of tumor cells (characteristic of lung, breast, ovarian,  
 CC prostate, pancreatic, gastric, colon, renal, bladder cancers; melanoma,  
 CC hepatoma, sarcoma and lymphoma) in a patient and for inhibiting tumor  
 CC growth. Y02125-32 represent peptide linkers used to make the  
 CC multifunctional proteins of the invention.  
 CC  
 CC Sequence 5 AA;  
 SO  
 Query Match 100.0%; Score 28; DB 20; Length 5;  
 Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
 Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 QY 1 GGGGS 5  
 Db 1 11111  
 1 999gs 5  
 RESULT 10  
 ID Y83210 standard; Peptide; 5 AA.  
 XX Y83210;  
 AC  
 XX 24-JUL-2000 (first entry)  
 DT  
 XX Peptide linker used in construction of a\_vb\_3 integrin/IFN alpha.  
 DE  
 XX Biconjugate; a\_vb\_3 integrin; interferon alpha; angiogenesis;  
 KW cancer; tumour; osteoporosis; Paget's disease; Kaposi's sarcoma;  
 KW periodontal disease; metastasis; neoplasia; retinopathy; arthritis;  
 KW psoriasis; leukaemia; malignant melanoma; atherosclerosis;  
 KW smooth muscle cell migration; inhibition; treatment; antagonist;  
 KW angina; thrombosis; restenosis; antiviral; antifungal;  
 KW antibacterial.  
 KW  
 XX Synthetic.  
 OS  
 XX WO200009143-A1.  
 PN  
 XX 24-FEB-2000.  
 PD  
 XX 07-APR-1999; 99WO-US04296.  
 PE  
 XX 13-AUG-1998; 98US-0096442.  
 PR  
 XX (SEAR ) SEARLE & CO G D.  
 PA  
 XX Fok KF, Tjoeng FS;  
 PI WPI: 2000-205894/18.  
 DR  
 XX New biconjugates comprising an avb3 antagonist and a  
 PT metastatic-associated receptor ligand, useful for treating cancer and  
 PT other angiogenic diseases, or as antiviral, antifungal or antibacterial  
 PT agents  
 PS Claim 19; Page 88; 123pp; English.  
 XX  
 CC Biconjugates comprising one or more a\_vb\_3 antagonist moieties  
 CC coupled to a peptide or polypeptide having anti-angiogenic properties.  
 CC can be used for treating a human patient with an  
 CC angiogenesis-mediated disease, e.g. cancer, arthritis, or macular  
 CC degeneration. The a\_vb\_3 integrin is normally associated with  
 CC endothelial cells but can promote the formation of blood vessels  
 CC (angiogenesis) in tumours. The a\_vb\_3 integrin is also known to

CC play a role in tumour metastasis, neoplasia, osteoporosis,  
CC Paget's disease, retinopathy, arthritis, periodontal disease,  
CC psoriasis and smooth muscle cell migration. Interferon alpha is a  
CC family of proteins which possess complex antiviral, antineoplastic  
CC and immunomodulating activities. Interferon alpha is effective  
CC against a variety of cancers including hairy cell leukaemia,  
CC chronic myelogenous leukaemia, malignant melanoma and Kaposi's  
CC sarcoma. Multi-functional bioconjugates comprising both a  $\gamma$ -b<sub>3</sub>  
CC antagonists and interferon alpha 2b can exhibit greater biological  
CC activity when compared to a single factor or having improved  
CC half-life or decreased adverse side effects, or a combination of  
CC these properties. They can be used for inhibiting elevated levels  
CC of tumor antigens, inhibiting the proliferation of tumor cells and  
CC inhibiting tumor growth. The bioconjugates can also be used for  
CC treating e.g. osteoporosis, humoral hypercalcemia of malignancy,  
CC Paget's disease, retinopathy including diabetic retinopathy,  
CC arthritis including rheumatoid arthritis, periodontal disease,  
CC psoriasis, thrombosis, angina, atherosclerosis, smooth muscle cell  
CC migration and restenosis in a mammal. They are also useful as  
CC antiviral, antifungal and antibacterial agents. This sequence is a  
CC peptide linker used in the construction of the multi-functional  
CC bioconjugates.

SQ Sequence 5 AA;

Query Match	100.0%;	Score 28;	DB 21;	Length 5;
Best Local Similarity	100.0%;	Pred. No. 2.1e+05;		
Matches	5;	Conservative	0;	Mismatches 0;
			Indels	0;
			Gaps	0

QY	1	GGGS	5
Db	1	ggggs	5

```

RESULT 11
Y43750
ID Y43750 standard; 'peptide; 5 AA

```

AC Y43750-

DT 11-FEB-2000 (first entry)

DE Linker used to construct a bispecific single-chain antibody.

bscCD19xCD3 antibody; bispecific single-chain fragment; CD19 antigen;

KW cytotoxic T-lymphocyte; B-cell malignancy; myasthenia gravis;

KW Hashimoto thyroiditis; Goodpasture syndrome; B-cell depletion;

OS Synthetic.

PN WO9954440-A1.

PD 28-OCT-1999.

PF 21-APR-1999; 99WO-EP02693.

PR 21-APR-1998; 98EP-0107269.

PA (DOER/) DOERKEN B.  
PA (RIET/) RIETHMUELLER G.

PI Kufer P, Lutterbuese R, Bargou R, Loeffler A;

DR WPI; 2000-013241/01.

PT Novel multifunctional polypeptide for treating B-cell malignancies  
PT especially non-Hodgkin Lymphoma -  
XX  
PS Claim 10; Page 49; 91pp; English.

XX The present sequence represents a linker used in the construction  
CC of bispecific single-chain polypeptides of the invention. These  
CC polypeptides comprise domains providing binding-site of immunoglobulin  
CC chains or antibodies specifically recognizing CD19 and CD3 antigen.  
CC The polypeptide destroys CD19-positive target cells without any need  
CC of T-cell pre and/or co-stimulation, by recruiting cytotoxic  
CC T-lymphocytes and so specific lysis by T-cells rather than a direct  
CC effect by an antibody is achieved. The bispecific single-chain  
CC polypeptides, or nucleotides encoding them, are used for the treatment  
CC of B-cell malignancies, B-cell mediated autoimmune diseases like  
CC myasthenia gravis, Morbus Basedow, Hashimoto thyroiditis or Goodpasture  
CC syndrome or for the depletion of B-cells and more particularly  
CC non-Hodgkin lymphoma in mammals preferably human. They can also delay  
CC the pathological conditions caused by these diseases, and can be used  
CC for detecting these diseases. The polynucleotide is used for gene  
CC therapy. The polypeptides are also used for identifying compounds  
CC modulating B-cell/T-cell mediated immune response with can in turn be  
CC used for treating cancer, its related diseases and also for inhibiting  
CC viral diseases by preventing viral infection.

SQ Sequence 5 AA;

Query Match	100.0%;	Score 28;	DB 21;	Length 5;
Best Local Similarity	100.0%;	Pred. No. 2.1e+05;		
Matches	5;	Conservative	0;	Mismatches 0;
			Indels	0;
			Gaps	0

QY	1	GGGS	5
Db	1	ggggs	5

RESULT	12
Y54917	
ID	Y54917 standard; peptide; 5 AA.

AC Y5491.7;

DT 14-FEB-2000 (first entry)

DE Linker from IL-12 fusion protein.

KW Interleukin-12; IL-12; fusion protein; IL-12 p35 subunit; B7 protein;

OS Synthetic

PN US5994104-A

PD 30-NOV-1999

PF 08-NOV-1996; 96US-0751767.

PR 08-NOV-1996; 96US-0751767.

PA (UNLO ) ROYAL FREE HOSPITAL SCHOOL MED.

PI Anderson RJ, Prentice HG, MacDonald ID,

PT Nucleic acid constructs encoding interleukin-12 fusion proteins useful for treating leukemia and other cancers -

CC This sequence represents a linker that can be used in an interleukin-12  
CC fusion protein. The invention relates to an isolated nucleic acid  
CC construct (1) comprising a region encoding an interleukin-12 (IL-12)  
CC fusion protein (comprising an IL-12 p35 subunit, an IL-12 p40 subunit and  
CC a linker peptide (joining the subunits)), and a region encoding a B7  
CC protein. (1) may be used to produce IL-12 fusion proteins according to

CC standard recombinant DNA methodologies. The fusion proteins may be  
CC produced either in vitro in a fermentation culture or in vivo as part of  
CC a gene therapy protocol (in this case (I) is used to transform a patients  
CC cells, which then secrete the functional polypeptide to supplement the  
CC patients own production of IL-12 or to rectify mutations which lead to  
CC the expression of inactive polypeptides). The fusion proteins produced in  
CC this way may be used to treat any disease which responds to IL-12 such as  
CC tumours (both solid and dispersed of the kidney, breast, colon, ovarian  
CC and cervical tumours and melanomas) and in particular, tumours of the  
CC blood such as leukaemia. Alternatively, the polypeptides may be used as  
CC antigens in the production of antibodies to IL-12 and to assay for  
CC agonists and antagonists of its activity. The antibodies and antagonists  
CC may be used to inhibit the activity of IL-12. (I) may also be used  
CC diagnostically as a probe which hybridizes to sequences encoding IL-12  
CC and the antibodies may be used to detect the presence of IL-12  
CC polypeptides in samples. They may be used diagnostically to quantitate  
CC the expression of the polypeptide by patients and hence which subjects  
CC may be in need of restorative therapy.  
CC  
CC  
SQ Sequence 5 AA;

Query Match 100.0%; Score 28; DB 21; Length 5;  
Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 1 GGGGS 5  
| | | | |  
DB 1 9999s 5

RESULT 13  
R62168  
ID R62168 standard; Protein: 6 AA.

AC R62168;

DT 03-MAY-1995 (first entry)

DE U1 snRNP 70K protein amino acids 5-10, homologous to HSV-1 IE motif.

Small ribonucleoprotein complex; U1 snRNP; 70K protein; epitope;  
antibody; immunoinfective cluster virus; nuclear protein antigen;  
systemic rheumatic disorder; herpes simplex virus; HSV-1;  
immediate early protein; systemic lupus erythematosus; scleroderma.

Homo sapiens.

WO9420141-A.

15-SEP-1994.

10-MAR-1994; 94WO-US02631.

11-MAR-1993; 93US-0029850.

(UWSC-) UNIV SOUTHERN CALIFORNIA.

Douvas A, Ehresmann G, Takehana Y;

WPI: 1994-302689/37.

Methods for treating immunoinfective cluster virus infections -  
utilise antibodies or fragments characteristic of auto antibodies  
produced by patients with rheumatic disorders  
Disclosure; Page 63; 106pp; English.

A comparison of the U1 snRNP 70K protein sequence with proteins  
from immunoinfective cluster viruses revealed widespread  
homologies. The importance of these homologous motifs is that they  
are epitopes for autoantibodies occurring in high titres in systemic  
rheumatic disorders. Sera from such patients could be used for

CC treatment of immunoinfective cluster virus infections.  
XX  
SQ Sequence 6 AA;

Query Match 100.0%; Score 28; DB 15; Length 6;  
Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
QY 1 GGGGS 5  
| | | | |  
DB 2 9999s 6

RESULT 14  
W17095  
ID W17095 standard; peptide: 6 AA.

AC W17095;

DT 14-SEP-1999 (first entry)

DE Gly(5)-Ser linker peptide for chimeric protein construct.

Haematopoietic protein; human; granulocyte-colony stimulating factor;  
G-CSF; interleukin; c-mpl ligand; linker; gene therapy; aplastic anaemia;  
stem cell expansion; leukaemia; neutropenia; vector; bone marrow;  
thrombocytopaenia; blood cell activation; growth.

Synthetic.

WO9712985-A2.

10-APR-1997.

04-OCT-1996; 96WO-US15774.

05-OCT-1995; 95US-0004834.

(SEAR) SEARLE & CO G.D.

Bauer SC, Baum CM, Caparon MH, Feng Y, Giri JG;  
Klein BK, Lee SC, McKearn JP, McWhorter CA, Statten NR;

Summers NL, Zurfluh L;

WPI: 1997-226228/20.

Multi-functional haematopoietic receptor agonists - used to  
stimulate the production of haematopoietic cells in patients

Disclosure; Page 33; 616pp; English.

The invention relates to a novel haematopoietic protein (HP) comprising  
an amino acid (AA) sequence of formula: R1-L1-R2; R2-L1-R1; R1-R2; or  
R2-R1; where R1 and R2 are independently selected from: (I) a modified  
human granulocyte-colony stimulating factor (hG-CSF) AA sequence;  
(II) a modified human interleukin-3 (hIL-3) AA sequence; (III) a modified  
human c-mpl ligand; and a colony stimulating factor (CSF); and L1 = a  
linker capable of linking R1 to R2. This sequence represents an example  
of a linker used to construct the proteins of the invention.  
Vectors comprising the nucleic acid molecules are useful for the  
recombinant production of HP. The nucleic acid molecules are useful in  
gene therapy. The HP's are useful for stimulating the production of  
haematopoietic cells in patients, selective ex vivo expansion of stem  
cells and for treatment of haematopoietic disorders. Disorders that  
can be treated include leukaemia, neutropenia, aplastic anaemia and  
thrombocytopaenia. In vitro uses include the ability to stimulate bone  
marrow and blood cell activation and growth before infusion into the  
patients.  
Sequence 6 AA;

Query Match 100.0%; Score 28; DB 18; Length 6;

Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

OY 1 GGGS 5  
11111  
DB 2 ggggs 6

## RESULT 15

W21967  
ID W21967 standard; peptide: 6 AA.  
XX

AC W21967;  
XX

DT 03-DEC-1997 (first entry)  
XX

DE Linker #1 for immunotoxin containing Pseudomonas exotoxin.  
XX

XX PCR; primer; amplify; polymerase chain reaction; antibody; immunotoxin;  
KW variable heavy chain; VH; murine monoclonal antibody; Lewis; carcinoma;  
KW carbohydrate antigen; Pseudomonas exotoxin; proteolytic activation;  
KW cytotoxic activity; tumour; autoimmune condition; rheumatoid arthritis;  
KW graft versus host disease; organ transplant rejection; type I diabetes;  
KW multiple sclerosis; systemic lupus erythematosus; myasthenia gravis;  
KW T cell; B cell; cytosol; bone marrow; transplant; therapy.  
XX

XX Synthetic.  
OS

XX MO9713529-A1.  
XX

XX 17-APR-1997.  
XX

XX 11-OCT-1996; 96WO-US16327.  
XX

XX 13-OCT-1995; 95US-0005388.  
XX

XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
XX

XX Kuan C, Pastan I;  
XX

XX WPI: 1997-235666/21.  
XX

XX Immunotoxin(s) comprising Pseudomonas exotoxin linked to  
PT disulphide stabilised variable heavy and light chain regions of an  
PT antibody - useful for killing target cells bearing characteristic  
PT marker  
XX

PS Claim 5, Page 49; 64pp; English.  
XX

XX W21967-W21969 represent linkers used in the immunotoxins of the  
CC invention. The immunotoxins bind to target cells, and comprise, a  
CC Pseudomonas exotoxin (PE) that does not need proteolytic activation for  
CC cytotoxic activity fused to a VH framework region of an Fv antibody (Ab)  
CC fragment. The VH chain region is bound through at least one disulphide  
CC bond to a variable light (VL) chain framework region. The PE is lacking  
CC residues 1-279 and is at least 10-fold more cytotoxic to the target cells  
CC than an immunotoxin comprising PE attached to a VH chain framework region  
CC of an Fv Ab fragment lacking a disulphide bond to a VL chain framework  
CC region. These sequences are used to join the VH chain region to the PE.  
CC The immunotoxins can be used for killing target cells in the treatment of  
CC tumours, autoimmune conditions, graft versus host disease, organ  
CC transplant rejection, type I diabetes, multiple sclerosis, rheumatoid  
CC arthritis, systemic lupus erythematosus, myasthenia gravis, etc, all  
CC caused by T and B cells. They can also be used to deliver an antibody to  
CC the cytosol of a cell, and in vitro in the elimination of harmful cells  
CC from bone marrow before transplant. The immunotoxins have high  
CC cytotoxicity to target cells and a small size to provide greater  
CC penetration to target cells.  
XX

XX Sequence 6 AA;  
SQ

Best Local Similarity 100.0%; Pred. No. 2.1e+05;  
Matches 5; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

OY 1 GGGS 5  
11111  
DB 2 ggggs 6

Search completed: March 15, 2001, 10:52:21  
Job time: 971 sec

Query Match 100.0%; Score 28; DB 18; Length 6;



